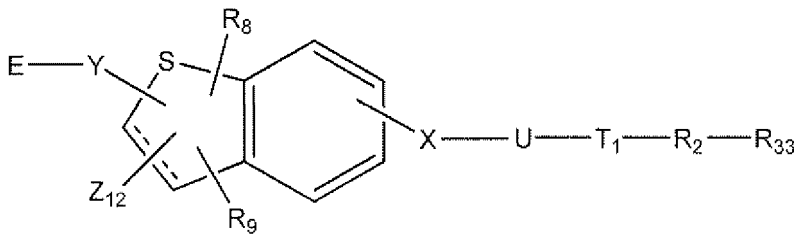
Amendments to the Claims

Please amend Claim 46. The Claim Listing below will replace all prior versions of the claims in the application:

Claim Listing

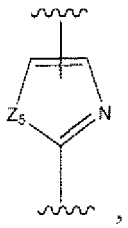
1-3. (Cancelled)

4. (Previously presented) A compound represented by the following Structural Formula:



or a stereoisomer, pharmaceutically acceptable salt, or hydrate thereof, wherein:

(a) T1 is a moiety of formula



,

optionally substituted with

one substituent selected from C_1-C_8 alkenyl, aryl-C_{0-4}-alkyl, aryl-C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, and C3-C6 cycloalkylaryl-C_{0-2}-alkyl, wherein C_1-C_8 alkyl, C_1-C_8 alkenyl, aryl-C_{0-4}-alkyl, aryl-C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, and C3-C6 cycloalkylaryl-C_{0-2}-alkyl are each optionally substituted with from one to three substituents independently selected from R1′; or

one substituent selected form C_3-C_6 alkyl, C_1-C_6 haloalkyl, and C_1-C_6 alkyloxo; and;

(b) R1′, R26, R27, R28, R31, Z14′, and Z15′ are each independently selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C_1-C_6 alkyl, C_1-C_6 alkyl-COOR12, C_1-C_6 alkoxy, C_1-C_6 haloalkyl, C_1-C_6 haloalkyloxy, C_3-C_7 cycloalkyl, aryloxy, aryl-C_{0-4}-alkyl, heteroaryl, heterocycloalkyl, C(O)R13, COOR14, OC(O)R15, $OS(O)_2R16$, $N(R17)_2$, NR18C(O)R19, $NR20SO_2R21$, SR22, S(O)R23, $S(O)_2R24$, and $S(O)_2N(R25)_2$; R12, R13, R14, R15, R16, R17, R18, R19, R20, R21, R22, R23, R24 and R25 are each independently selected from the group consisting of hydrogen, C_1-C_6 alkyl and aryl;

(c) R2 is selected from the group consisting of C_0-C_8 alkyl and C_{1-6}-heteroalkyl;

(d) X is selected from the group consisting of a bond, O, S, $S(O)_2$ and N;

(e) U is an aliphatic linker wherein one carbon atom of the aliphatic linker may be replaced with O, NH or S, and wherein such aliphatic linker is optionally substituted with R30;

(f) Y is selected from the group consisting of C, O, S, NH and a single bond;

(g) E is C(R3)(R4)A or A and wherein

(i) A is selected from the group consisting of carboxyl, tetrazole, C_1-C_6 alkylnitrile, carboxamide, sulfonamide and acylsulfonamide; wherein sulfonamide, acylsulfonamide and tetrazole are each optionally substituted with from one to two groups independently selected from R^7;

(ii) each R^7 is independently selected from the group consisting of hydrogen, C_1-C_6 haloalkyl, aryl-C_0-C_4 alkyl and C_1-C_6 alkyl, wherein such alkyl and arylalkyl are each optionally substituted with from one to two groups independently selected from R7′; each R7′ is independently selected from halo, C_1-C_6 alkyl, and haloC_1-C_6 alkyl;

(iii) R3 is selected from the group consisting of hydrogen, C_1-C_5 alkyl, and C_1-C_5 alkoxy; and

(iv) R4 is selected from the group consisting of hydrogen, C_1-C_5 alkyl, C_1-C_5 alkoxy, aryloxy, C_3-C_6 cycloalkyl, and aryl C_0-C_4 alkyl, and R3 and R4 are optionally combined to form a C_3-C_4 cycloalkyl, and wherein alkyl, alkoxy,

cycloalkyl and aryl-alkyl are each optionally substituted with one to three each independently selected from R26;

(h) Z5 is S or O;

(i) Z12 is selected from the group consisting of hydrogen and -Z13C_0–C_3alkylZ14;

(j) Z13 is selected from the group consisting of a single bond, CO, CO_2, CONZ15, and SO_2;

(k) Z14 is selected from the group consisting of aryl and heteroaryl, wherein the aryl and heteroaryl is each optionally substituted with from one to three substituents independently selected from Z14′;

(l) Z15 is selected from the group consisting of hydrogen, aryl, and heteroaryl, wherein the aryl and heteroaryl is each optionally substituted with from one to three substituents independently selected from Z15′;

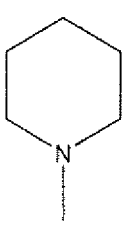
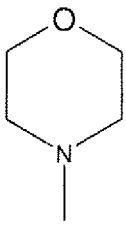
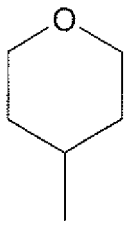
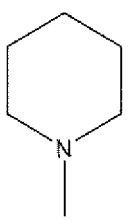
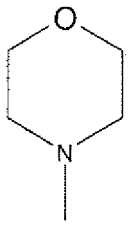
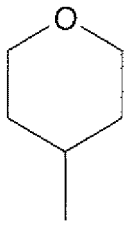
(m) R8 is selected from the group consisting of hydrogen, C_1-C_4 alkyl, C_1-C_4 alkylenyl, oxo, sulfo, and halo;

(n) R9 is selected from the group consisting of hydrogen, C_1-C_4 alkyl, C_1-C_4 alkylenyl, halo, aryl-C_0-C_4 alkyl, heteroaryl, C_1-C_6 allyl, oxo, sulfo, and OR29, and R8 and R9 together optionally combine to form a fused C5-C6 ring with the carbons to which they are attached, and wherein aryl-C_0-C_4 alkyl, heteroaryl are each optionally substituted with from one to three independently selected from R27; R29 is selected from the group consisting of hydrogen and C_1-C_4 alkyl;

(o) R10, R11 are each independently selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C_1-C_6 alkyl, C_0-C_6 alkyl-COOR12″, C_1-C_6 alkoxy, C_1-C_6 haloalkyl, C_1-C_6 haloalkyloxy, C_3-C_7 cycloalkyl, aryl-C_{0-4}-alkyl, aryl-C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, C3-C6 cycloalkylaryl-C_{0-2}-alkyl, aryloxy, C(O)R13′, COOR14′, OC(O)R15′, $OS(O)_2$R16′, $N(R17')_2$, NR18′C(O)R19′, NR20′SO_2R21′, SR22′, S(O)R23′, $S(O)_2$R24′, and $S(O)_2N(R25')_2$; and wherein aryl-C_{0-4}-alkyl, aryl- C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, and C3-C6 cycloalkylaryl-C_{0-2}-alkyl are each optionally substituted with from one to three independently selected from R28;

(p) R12′, R12″, R13′, R14′, R15′, R16′, R17′, R18′, R19′, R20′, R21′, R22′, R23′, R24′, and R25′ are each independently selected from the group consisting of hydrogen, C_1-C_6 alkyl and aryl;

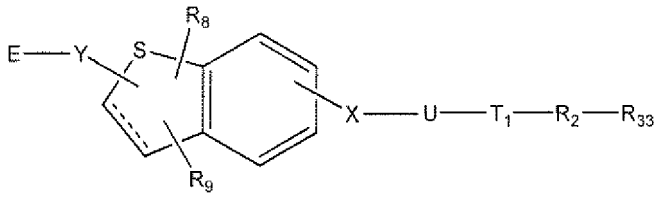
(q) R30 is selected from the group consisting of C_1-C_6 alkyl, aryl-C_{0-4}-alkyl, aryl-C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, and C3-C6 cycloalkylaryl-C_{0-2}-alkyl, and wherein C_1-C_6 alkyl, aryl-C_{0-4}-alkyl, aryl- C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, and C3-C6 cycloalkylaryl-C_{0-2}-alkyl are each optionally substituted with from one to three substituents each independently selected from R31;

(r) R33 is selected from the group consisting of phenyl, thiophene, pyridine, piperidine,  ,  , and  , wherein the phenyl, thiophene, pyridine, piperidine,  ,  , and  , are each optionally substituted with R10 and R11;

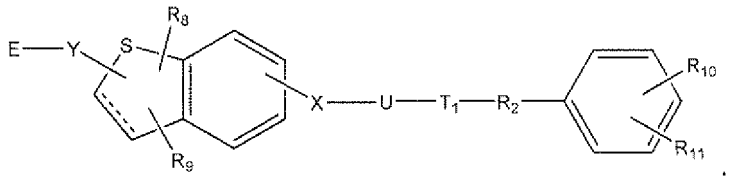
wherein "----" are each independently an optional bond to form a double bond at the indicated position.

5. (Cancelled)

6. (Previously Presented) The compound of Claim 4, wherein the compound is represented by the following Structural Formula:

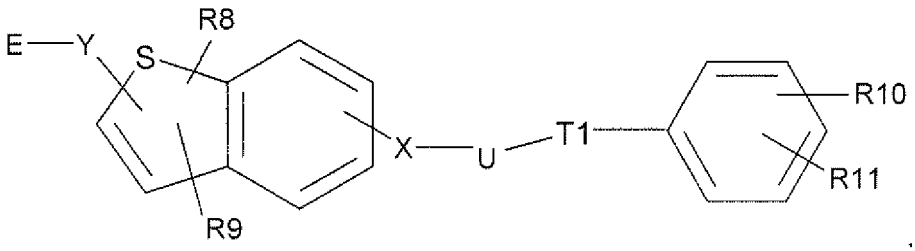


7. (Previously Presented) The compound of Claim 6, wherein the compound is represented by the following Structural Formula:



8-9. (Canceled)

10. (Previously Presented) The compound of Claim 7 wherein the compound is represented by the following Structural Formula:



11. (Previously presented) The compound of Claim 10 wherein:

X is -O-;

E is $C(R3)(R4)CO_2H$ or CO_2H;

R1, R3, and R4 are each independently selected from the group consisting of hydrogen and C_1-C_2 alkyl;

R10 and R11 are each independently selected from the group consisting of hydrogen, halo, oxo, C_1-C_6 alkyl, C_1-C_6 alkyl-COOR12", C_1-C_6 alkoxy, C_1-C_6 haloalkyl, and C_1-C_6 haloalkyloxy; and

U is saturated C_1-C_3 alkyl optionally substituted with C_1-C_3 alkyl.

12-45. (Cancelled)

46. (Currently Amended) A compound selected from the group consisting of:
{6-[5-Methyl-2-(4-trifluoromethyl-phenyl)-oxazol-4-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid;
{4-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-thiazol-5-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid;
{4-[5-Methyl-2-(4-trifluoromethyl-phenyl)-oxazol-4-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid;
(6-{2-[5-Methyl-2-(4-trifluoromethyl-phenyl)-thiazol-4-yl]-propoxy}-benzo[b]thiophen-3-yl)-acetic acid;
(6-{1-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-thiazol-5-yl]-ethoxy}-benzo[b]thiophen-3-yl)-acetic acid;
(6-{2-[5-Methyl-2-(4-trifluoromethyl-phenyl)-oxazol-4-yl]-ethoxy}-benzo[b]thiophen-3-yl)-acetic acid;
(6-{2-[5-Methyl-2-(4-trifluoromethyl-phenyl)-oxazol-4-yl]-propoxy}-benzo[b]thiophen-3-yl)-acetic acid;
(6-{2-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-thiazol-5-yl]-propoxy}-benzo[b]thiophen-3-yl)-acetic acid;
(*R*)-(6-{2-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-thiazol-5-yl]-propoxy}-benzo[b]thiophen-3-yl)-acetic acid;
(*S*)-(6-{2-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-thiazol-5-yl]-propoxy}-benzo[b]thiophen-3-yl)-acetic acid;
(*R*)-(4-{2-[5-Methyl-2-(4-trifluoromethyl-phenyl)-thiazol-4-yl]-propoxy}-benzo[b]thiophen-3-yl)-acetic acid;

(*S*)-(4-{2-[5-Methyl-2-(4-trifluoromethyl-phenyl)-thiazol-4-yl]-propoxy}-benzo[b]thiophen-3-yl)-acetic acid;

(4-{2-[5-Methyl-2-(4-trifluoromethyl-phenyl)-thiazol-4-yl]-ethoxy}-benzo[b]thiophen-3-yl)-acetic acid;

Racemic-(4-{1-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-thiazol-5-yl]-ethoxy}-benzo[b]thiophen-3-yl)-acetic acid;

~~(6-{2-[5-Methyl-2-(4-trifluoromethyl-phenyl)-oxazol-4-yl]-propoxy}-benzofuran-3-yl)-acetic acid;~~

~~(6-{2-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-thiazol-5-yl]-propoxy}-benzofuran-3-yl)-acetic acid;~~

~~(6-{2-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-oxazol-5-yl]-propoxy}-benzofuran-3-yl)-acetic acid;~~

~~(6-{2-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-oxazol-5-yl]-ethoxy}-benzofuran-3-yl)-acetic acid;~~

~~{6-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-oxazol-5-ylmethoxy]-benzofuran-3-yl}-acetic acid;~~

~~(6-{1-Methyl-1-[4-methyl-2-(4-trifluoromethyl-phenyl)-oxazol-5-yl]-ethylsulfanyl}-benzofuran-3-yl)-acetic acid;~~

~~{6-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-oxazol-5-ylmethylsulfanyl]-benzofuran-3-yl}-acetic acid;~~

~~(6-{2-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-oxazol-5-yl]-ethylsulfanyl}-benzofuran-3-yl)-acetic acid;~~

~~(6-{1-[4-Methyl-2-(4-trifluoromethyl-phenyl)-oxazol-5-yl]-ethylsulfanyl}-benzofuran-3-yl)-acetic acid;~~

~~2-{6-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-oxazol-5-ylmethoxy]-benzofuran-3-yl}-propionic acid;~~

~~2-(6-{2-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-oxazol-5-yl]-ethoxy}-benzofuran-3-yl)-propionic acid;~~

~~(6-{2-[5-Methyl-2-(4-trifluoromethyl-phenyl)-thiazol-4-yl]-ethoxy}-benzofuran-3-yl)-acetic acid;~~

~~(*R*)-(6-{2-[5-Methyl-2-(4-trifluoromethyl-phenyl)-thiazol-4-yl]-propoxy}-benzofuran-3-yl)-acetic acid (Isomer 2);~~

~~(*S*)-(6-{2-[5-Methyl-2-(4-trifluoromethyl-phenyl)-thiazol-4-yl]-propoxy}-benzofuran-3-yl)-acetic acid;~~

~~{5-[2-(5-Methyl-3-phenyl-pyrazol-1-yl)-ethoxy]-indol-1-yl}-acetic acid;~~

~~(1-Methyl-6-{2-[3-methyl-1-(4-trifluoromethyl-phenyl)-1H-pyrazol-4-yl]-propoxy}-1H-indol-3-yl)-acetic acid;~~

~~{5-[5-(4-Trifluoromethyl-phenyl)-thiophen-2-ylmethoxy]-indol-1-yl}-acetic acid;~~

~~(5-{2-[3-Methyl-5-(4-trifluoromethyl-phenyl)-thiophen-2-yl]-propoxy}-indol-1-yl)-acetic acid;~~

{6-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-thiazol-5-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid; and

2-(6-((4-tert-butyl-2-(4-(trifluoromethyl)phenyl)thiazol-5-yl)methylthio)-benzo[*b*]thiophen-3-yl)acetic acid.

47-49. (Cancelled)

50. (Previously Presented) A method of treating a mammal in need of treatment for a disease, wherein the disease is treatable by activating a peroxisome proliferator activated receptor, comprising the step of administering to the mammal in need thereof a therapeutically effective amount of the compound of Claim 4, wherein the disease is selected from diabetes mellitus, Syndrome X, and atherosclerosis.

51-53. (Cancelled)

54. (Previously Presented) The method of Claim 50, wherein the disease is diabetes mellitus.

55. (Previously Presented) The method of Claim 50, wherein the disease is Syndrome X.

56-100. (Cancelled)

101. (Previously Presented) A compound, wherein the compound is:
{6-[4-Isopropyl-2-(4-trifluoromethyl-phenyl)-thiazol-5-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid; or
2-(6-((4-tert-butyl-2-(4-(trifluoromethyl)phenyl)thiazol-5-yl)methylthio)benzo[*b*]thiophen-3-yl)acetic acid;
or a stereoisomer, pharmaceutically acceptable salt, or hydrate thereof.

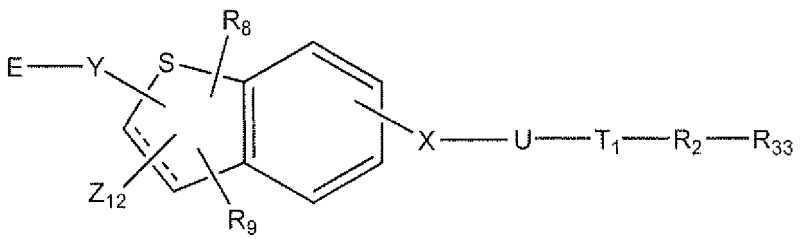
102-107. (Cancelled)

108. (Previously Presented) A method of treating a human subject in need of treatment for a disease selected from the group consisting of diabetes mellitus, Syndrome X, and atherosclerosis, comprising the step of administering to the subject in need thereof a therapeutically effective amount of the compound of Claim 101.

109-114. (Cancelled)

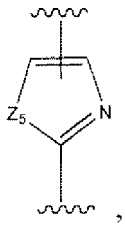
115. (Previously Presented) A method of treating a mammal in need of treatment for a disease, wherein the disease is treatable by activating a peroxisome proliferator activated receptor, comprising the step of administering to the mammal in need thereof a therapeutically effective amount of the compound of Claim 46, disease selected from the group consisting of diabetes mellitus, Syndrome X, and atherosclerosis.

116. (Previously Presented) A compound represented by the following Structural Formula:



or a stereoisomer, pharmaceutically acceptable salt, or hydrate thereof, wherein:

(s) T1 is a moiety of formula



optionally substituted with

one substituent selected from C_1-C_8 alkenyl, aryl-C_{0-4}-alkyl, aryl-C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, and C3-C6 cycloalkylaryl-C_{0-2}-alkyl, wherein C_1-C_8 alkyl, C_1-C_8 alkenyl, aryl-C_{0-4}-alkyl, aryl-C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, and C3-C6 cycloalkylaryl-C_{0-2}-alkyl are each optionally substituted with from one to three substituents independently selected from R1′; or

one substituent selected form C_1-C_6 haloalkyl, and C_1-C_6 alkyloxo; and;

(t) R1′, R26, R27, R28, R31, Z14′, and Z15′ are each independently selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C_1-C_6 alkyl, C_1-C_6 alkyl-COOR12, C_1-C_6 alkoxy, C_1-C_6 haloalkyl, C_1-C_6 haloalkyloxy, C_3-C_7 cycloalkyl, aryloxy, aryl-C_{0-4}-alkyl, heteroaryl, heterocycloalkyl, C(O)R13, COOR14, OC(O)R15, $OS(O)_2R16$, $N(R17)_2$, NR18C(O)R19, $NR20SO_2R21$, SR22, S(O)R23, $S(O)_2R24$, and $S(O)_2N(R25)_2$; R12, R13, R14, R15, R16, R17, R18, R19, R20, R21, R22, R23, R24 and R25 are each independently selected from the group consisting of hydrogen, C_1-C_6 alkyl and aryl;

(u) R2 is selected from the group consisting of C_0-C_8 alkyl and C_{1-6}-heteroalkyl;

(v) X is selected from the group consisting of a bond, O, S, $S(O)_2$ and N;

(w) U is an aliphatic linker wherein one carbon atom of the aliphatic linker may be replaced with O, NH or S, and wherein such aliphatic linker is optionally substituted with R30;

(x) Y is selected from the group consisting of C, O, S, NH and a single bond;

(y) E is C(R3)(R4)A or A and wherein

(i) A is selected from the group consisting of carboxyl, tetrazole, C_1-C_6 alkylnitrile, carboxamide, sulfonamide and acylsulfonamide; wherein

sulfonamide, acylsulfonamide and tetrazole are each optionally substituted with from one to two groups independently selected from R^7;

(ii) each R^7 is independently selected from the group consisting of hydrogen, C_1-C_6 haloalkyl, aryl-C_0-C_4 alkyl and C_1-C_6 alkyl, wherein such alkyl and arylalkyl are each optionally substituted with from one to two groups independently selected from R7'; each R7' is independently selected from halo, C_1-C_6 alkyl, and haloC_1-C_6 alkyl;

(iii) R3 is selected from the group consisting of hydrogen, C_1-C_5 alkyl, and C_1-C_5 alkoxy; and

(iv) R4 is selected from the group consisting of hydrogen, C_1-C_5 alkyl, C_1-C_5 alkoxy, aryloxy, C_3-C_6 cycloalkyl, and aryl C_0-C_4 alkyl, and R3 and R4 are optionally combined to form a C_3-C_4 cycloalkyl, and wherein alkyl, alkoxy, cycloalkyl and aryl-alkyl are each optionally substituted with one to three each independently selected from R26;

(z) Z5 is S or O;

(aa) Z12 is selected from the group consisting of hydrogen and -Z13C_0–C_3alkylZ14;

(bb) Z13 is selected from the group consisting of a single bond, CO, CO_2, CONZ15, and SO_2;

(cc) Z14 is selected from the group consisting of aryl and heteroaryl, wherein the aryl and heteroaryl is each optionally substituted with from one to three substituents independently selected from Z14';

(dd) Z15 is selected from the group consisting of hydrogen, aryl, and heteroaryl, wherein the aryl and heteroaryl is each optionally substituted with from one to three substituents independently selected from Z15';

(ee) R8 is selected from the group consisting of hydrogen, C_1-C_4 alkyl, C_1-C_4 alkylenyl, oxo, sulfo, and halo;

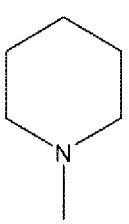
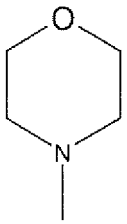
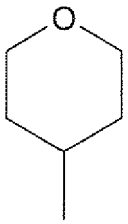
(ff) R9 is selected from the group consisting of hydrogen, C_1-C_4 alkyl, C_1-C_4 alkylenyl, halo, aryl-C_0-C_4 alkyl, heteroaryl, C_1-C_6 allyl, oxo, sulfo, and OR29, and R8 and R9 together optionally combine to form a fused C5-C6 ring with the

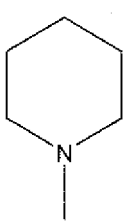
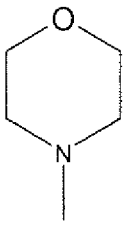
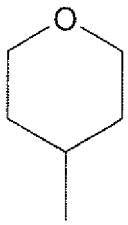
carbons to which they are attached, and wherein aryl-C_0-C_4 alkyl, heteroaryl are each optionally substituted with from one to three independently selected from R27; R29 is selected from the group consisting of hydrogen and C_1-C_4 alkyl;

(gg) R10, R11 are each independently selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C_1-C_6 alkyl, C_0-C_6 alkyl-COOR12″, C_1-C_6 alkoxy, C_1-C_6 haloalkyl, C_1-C_6 haloalkyloxy, C_3-C_7 cycloalkyl, aryl-C_{0-4}-alkyl, aryl-C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, C3-C6 cycloalkylaryl-C_{0-2}-alkyl, aryloxy, C(O)R13′, COOR14′, OC(O)R15′, $OS(O)_2R16'$, $N(R17')_2$, NR18′C(O)R19′, $NR20'SO_2R21'$, SR22′, S(O)R23′, $S(O)_2R24'$, and $S(O)_2N(R25')_2$; and wherein aryl-C_{0-4}-alkyl, aryl- C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, and C3-C6 cycloalkylaryl-C_{0-2}-alkyl are each optionally substituted with from one to three independently selected from R28;

(hh) R12′, R12″, R13′, R14′, R15′, R16′, R17′, R18′, R19′, R20′, R21′, R22′, R23′, R24′, and R25′ are each independently selected from the group consisting of hydrogen, C_1-C_6 alkyl and aryl;

(ii) R30 is selected from the group consisting of C_1-C_6 alkyl, aryl-C_{0-4}-alkyl, aryl-C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, and C3-C6 cycloalkylaryl-C_{0-2}-alkyl, and wherein C_1-C_6 alkyl, aryl-C_{0-4}-alkyl, aryl- C_{1-6}-heteroalkyl, heteroaryl-C_{0-4}-alkyl, and C3-C6 cycloalkylaryl-C_{0-2}-alkyl are each optionally substituted with from one to three substituents each independently selected from R31;

(jj) R33 is selected from the group consisting of phenyl, thiophene, pyridine, piperidine,  ,  , and  , wherein the phenyl,

thiophene, pyridine, piperidine,  ,  , and  , are each optionally substituted with R10 and R11;

wherein "----" are each independently an optional bond to form a double bond at the indicated position.